PENDING CLAIMS 48-100 & 102-182 STAVRIANOPOULOS ET AL., U.S. PATENT APPLICATION SERIAL NO. 08/486,070 FILED JUNE 7, 1995

48. A composition of matter comprising:

a transparent non-porous or translucent non-porous system containing a fluid or solution, which system comprises:

- (i) a solid support; and
- (ii) a double-stranded oligonucleotide or polynucleotide which is directly or indirectly fixed or immobilized to said solid support wherein one of the strands produces a soluble signal generated or generatable from a chemical label or labels which comprise a signalling moiety or moieties.
- 49. The composition according to claim 48, wherein said solid support is contained within the transparent non-porous or translucent non-porous system.
- 50. The composition according to claim 48, wherein said solid support is porous or non-porous.
- 51. The composition according to claim 50, wherein said porous solid support comprises a porous polymeric material.
- 52. The composition according to claim 51, wherein said porous polymeric material is selected from the group consisting of dextran, cellulose and nitrocellulose.
- 53. The composition according to claim 50, wherein said non-porous solid support is selected from the group consisting of siliceous matter and non-porous polymeric material.
- 54. The composition according to claim 53, wehrein said siliceous matter comprises glass or a glass-coated surface.
- 55. The composition according to claim 133, wherein said plastic or plasticcoated surface is selected from the group consisting of polyethylene, polypropylene, polystyrene and epoxy.

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- 56. The composition according to claim 48, wherein said system is selected from the group consisting of a well, a tube, a cuvette and an apparatus that comprises a plurality of said wells, tubes or cuvettes.
- 57. The composition according to claim 56, wherein said well comprises a microtiter well.
- 58. The composition according to claim 56, wherein said wells in the apparatus comprise microtiter wells.
- 59. The composition according to claim 48, wherein said solid support and said system are composed of the same materials.
- 60. The composition according to claim 48, wherein said solid support and said system are composed of different materials.
- 61. The composition according to claim 48, wherein one of said oligonucleotide or polynucleotide strands is directly or indirectly fixed or immobilized to the solid support.
- 62. The composition according to claim 61, wherein said oligonucleotide or polynucleotide strand is directly fixed or immobilized to the solid support by sandwich hybridization.
- 63. The composition according to claim 48, wherein said double-stranded oligonucleotide or polynucleotide is selected from the group consisting of DNA, RNA and a DNA-RNA hybrid, or a combination of any of the foregoing.
- 64. The composition according to claim 48, wherein said label or labels are the signalling moiety or moieties.
- 65. The composition according to claims 48 or 64, wherein said label or labels are directly attached to the oligonucleotide or polynucleotide.

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- The composition according to claims 48 or 64, wherein said label or labels 66. are indirectly attached to the oligonucleotide or polynucleotide.
- The composition according to claim 66, wherein said label or labels are 67. indirectly attached to the oligonucleotide or polynucleotide through the formation of a complex.
- The composition according to claim 67, wherein said complex is selected 68. from the group consisting of biotin and avidin, biotin and streptavidin, a sugar and lectin, and an antigen and an antibody.
- The composition according to claim 66, wherein said label or labels are 69. indirectly attached to the oligonucleotide or polynucleotide through a bridging moiety.
- The composition according to claim 48, wherein the signalling moiety or 70. moieties of said label or labels are directly or indirectly attached thereto.
- The composition according to claim 48, wherein said signalling moiety or moieties are selected from the group consisting of an enzyme, a co-enzyme, a chelating agent, a chromagen, a fluorescent agent and a chemiluminescent agent.
- The composition according to claim 48, wherein said soluble signal is 72. generatable from a chromagen, or by fluorescence or chemiluminescence.
- The composition according to claim 72, wherein said soluble signal is 73. indirectly generatable by an enzyme or enzymatic reaction.
- The composition according to claim 48, wherein said soluble signal is 74. detectable by a technique selected from the group consisting of photometric techniques and colorimetric techniques.
- The composition according to claim 74, wherein said photometric techniques 75. comprise spectrophotometric techniques.

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- 76. The composition according to claim 48, wherein said soluble signal is selected from the group consisting of a colored product, a chemiluminescent product and a fluorescent product.
- 77. (Amended) A composition of matter comprising:

a transparent non-porous or translucent non-porous system containing a fluid or solution, which system comprises:

a double-stranded oligonucleotide or polynucleotide which is directly or indirectly fixed or immobilized to said system wherein one of the strands produces a soluble signal generated or generatable from a chemical label or labels which comprise a signalling moiety or moieties.

- 78. The composition according to claim 77, wherein said non-porous system is selected from the group consisting of siliceous matter and non-porous polymeric material.
- 79. The composition according to claim 78, wherein said siliceous matter comprises glass or a glass-coated surface.
- 80. The composition according to claim 134, wherein said plastic or plastic-coated surface is selected from the group consisting of polyethylene, polypropylene, polystyrene and epoxy.
- 81. The composition according to claim 77, wherein said system is selected from the group consisting of a well, a tube, a cuvette and an apparatus that comprises a plurality of said wells, tubes or cuvettes.
- 82. The composition according to claim 81, wherein said well comprises a microtiter well.
- 83. The composition according to claim 81, wherein said wells in the apparatus comprise microtiter wells.

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- 84. The composition according to claim 77, wherein one of said oligonucleotide or polynucleotide strands is directly or indirectly fixed or immobilized ot said system.
- 85. The composition according to claim 84, wherein said oligonucleotide or polynucleotide strand is directly fixed or immobilized to the system by sandwich hybridization.
- 86. The composition according to claim 48, wherein said double-stranded oligonucleotide or polynucleotide is selected from the group consisting of DNA, RNA and a DNA-RNA hybrid, or a combination of any of the foregoing.
- 87. The composition according to claim 77, wherein said label or lables are the signalling moiety or moieties.
- 88. The composition according to claims 77 or 87, wherein said label or labels are directly attached to the oligonucleotide or polynucleotide.
- 89. The composition according to claims 77 or 87, wherein said label or labels are indirectly attached to the oligonucleotide or polynucleotide.
- 90. The composition according to claim 89, wherein said label or labels are indirectly attached to the oligonucleotide or polynucleotide through the formation of a complex.
- 91. The composition according to claim 90, wherein said complex is selected from the group consisting of biotin and avidin, biotin and streptavidin, a sugar and a lectin, and an antigen and an antibody.
- 92. The composition according to claim 89, wherein said label or labels are indirectly attached to the oligonucleotide or polynucleotide through a bridging moiety.
- 93. The composition according to claim 77, wherein the signalling moiety or moieties of said label or labels are directly or indirectly attached thereto.

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- 94. The composition according to claim 77, wherein said signalling moiety or moieties are selected from the group consisting of an enzyme, a co-enzyme, a chelating agent, a chromagen, a fluorescent agent and a chemiluminescent agent.
- 95. The composition according to claim 77, wherein said soluble signal is generatable from a chromagen, or by fluorescence or chemiluminescence.
- 96. The composition according to claim 95, wherein said soluble signal is indirectly generatable by an enzyme or enzymatic reaction.
- 97. The composition according to claim 77, wherein said soluble signal is detectable by a technique selected from the group consisting of photometric techniques and colormetric techniques.
- 98. The composition according to claim 97, wherein said photometric techniques comprise spectrophotometric techniques.
- 99. The composition according to claim 77, wherein said soluble signal is selected from the group consisting of a colored product, a chemiluminescent and a fluorescent product.
- 100. An apparatus comprising:
- 1) one or more solution containing means, each comprising a transparent non-porous or translucent non-porous device;
- 2) means for forming a fixed or immobilized double-stranded oligonucleotide or polynucleotide hybrid to a solid support in said device, said hybrid comprising a chemical label or labels attached to one strand of said hybrid, said label or labels comprising a signalling moiety or moieties which are capable of generating a soluble signal; and
- 3) means for producing a quantifiable or measurable soluble signal generatable or generated from said chemical label or labels which comprise said signalling moiety or moieties.

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- 102. A transparent non-porous or translucent non-porous system containing a fluid or solution, which system comprises:
- (ii) an oligonucleotide or polynucleotide hybridized or hybridizable to an oligo-polynucleotide sequence, said oligonucleotide or polynucleotide in double-stranded form producing a soluble signal generated or generatable from a chemical label or labels which comprise a signalling moiety or moieties; and
- (ii) a solid support having directly or indirectly fixed or immobilized thereto said oligo- or polynucleotide sequence or said oligonucleotide or polynucleotide (i).
- 103. The system according to claim 102, wherein said solid support is contained within said transparent non-porous or translucent non-porous system.
- 104. The system according to claim 102, wherein the solid support is porous or non-porous.
- 105. The system according to claim 104, wherein said porous solid support comprises a porous polymeric material.
- 106. The system according to claim 105, wherein said porous polymeric material is selected from the group consisting of dextran, cellulose and nitrocellulose.
- 107. The system according to claim 104, wherein said non-porous solid support is selected from the group consisting of siliceous material and non-porous polymeric material.
- 108. The system according to claim 104, wherein said siliceous material comprises glass or a glass-coated surface.
- 109. The composition according to claim 135, wherein said plastic or plastic-coated surface is selected from the group consisting of polyethylene, polypropylene, polystyrene and epoxy.

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- 110. The system according to claim 102, wherein said system is selected from the group consisting of a well, a tube, a cuvette and an apparatus that comprises a plurality of wells or microtitre wells, tubes or cuvettes.
- 111. The system according to claim 110, wherein said well comprises a microtiter well.
- 112. The system according to claim 110, wherein said wells in the apparatus comprise microtiter wells.
- 113. The system according to claim 102, wherein said solid support and said system are composed of the same materials.
- 114. The system according to claim 102, wherein aid solid support and said system are composed of different materials.
- 115. The system according to claim 102, wherein said solid support is capable of indirectly fixing or immobilizing the oligo- or polynucleotide sequence or said oligonucleotide or polynucleotide (i).
- 116. The system according to claim 115, wherein said oligo-or polynucleotide sequence or said oligonucleotide or polynucleotide is indirectly fixed or immobilized to the solid support through the hybridization of a complementary oligo- or polynucleotide sequence.
- 117. The system according to claim 102, wherein said solid support is capable of indirectly fixing or immobilizing the oligonucleotide or polynucleotide.
- 118. The composition according to claim 102, wherein said double-stranded oligonucleotide or polynucleotide (i) or said oligo- or polynucleotide is selected from the group consisting of DNA, RNA and a DNA-RNA hybrid, or a combination of any of the foregoing.
- 119. The system according to claim 102, wherein said label or labels are the signaling moiety.

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- 120. The system according to claims 102 or 119, wherein said label or labels are directly attached to the oligonucleotide or polynucleotide.
- 121. The system according to claims 102 or 119, wherein said label or labels are indirectly attached to the oligonucleotide or polynucleotide.
- 122. The system according to claim 121, wherein said label or labels are indirectly attached to the oligonucleotide or polynucleotide through the formation of a complex.
- 123. The system according to claim 122, wherein said complex is selected from the group consisting of biotin and avidin, biotin and streptavidin, a sugar and a lectin, and an antigen and an antibody.
- 124. The system according to claim 121, wherein said label or labels are indirectly attached to the oligonucleotide or polynucleotide through a bridging moiety.
- 125. The system according to claim 102, wherein the signalling moiety or moieties of said label or labels are directly or indirectly attached thereto.
- 126. The system according to claim 102, wherein said signalling moiety or moieties are selected from the group consisting of an enzyme, a co-enzyme, a chelating agent, a chromagen, a fluorescent agent and a chemiluminescent agent.
- 127. The system according to calim 102, wherein said soluble signal is generatable from a chromagen, or by fluorescence or chemiluminescence.
- 128. The system according to claim 102, wherein said soluble signal is indirectly generatable by an enzyme or enzymatic reaction.
- 129. The system according to claim 102, wherein said soluble signal is detectable by a technique selected from the group consisting of photometric techniques and colorimetric techniques.

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- 130. The system according to claim 129, wherein said photometric techniques comprise spectrophotometric techniques.
- 131. The system according to claim 102, wherein said soluble signal is selected from the group consisting of a colored product, a chemiluminescent product and a fluorescent product.
- 132. An apparatus comprising:
 - 1) means for retaining or containing a fluid or solution;
- one or more transparent non-porous or translucent non-porous devices,
 each comprising a solid support;
- 3) means for forming a fixed or immobilized oligonucleotide or polynucleotide hybrid to said solid support, said hybrid comprising a chemical label or labels attached to said hybrid, said label or labels further comprising a signalling moiety or moieties capable of generating a soluble signal;
- 4) means for quantifying or detecting a soluble signal generatable or generated from said chemical label or labels comprising said signalling moiety or moieties; and
 - 5) fluid or solution.
- 133. The composition of claims 54 or 79, wherein said glass or glass-coated surface comprises porous glass.
- 134. The system of claim 108, wherein said glass or glass-coated surface comprises porous glass.
- 135. The composition of claim 53, wherein said non-porous polymeric material comprises plastic or a plastic-coated surface.
- 136. The composition of claim 78, wherein said non-porous polymeric material comprises plastic or a plastic-coated surface.
- 137. The composition of claim 107, wherein said non-porous polymeric material comprises plastic or a plastic-coated surface.

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- 138. The composition of claim 48, wherein said system is selected from the group consisting of a well, a tube, a cuvette and an apparatus that comprises a plurality of said wells, tubes or cuvettes, and said solid support is selected from the group consisting of dextran, cellulose, nitrocellulose, glass or a glass-coated surface and plastic or a plastic-coated surface.
- 139. The system of claim 102, wherein said system is selected from the group consisting of a well, a tube, a cuvette and an apparatus that comprises a plurality of said wells, tubes or cuvettes, and said solid support is selected from the group consisting of dextran, cellulose, nitrocellulose, glass or a glass-coated surface and plastic or a plastic-coated surface.
- 140. The composition of claim 48, wherein said system functions as the solid support.
- 141. The composition of claim 77, wherein said system functions as a solid support.
- 142. The system of claim 102, wherein said system functions as the solid support.
- 143. An array of substrate surfaces, said array comprising a plurality of nucleic acid strands fixed or immobilized to said substrate surfaces.
- 144. The array of claim 143, wherein each said substrate surfaces has been treated to enhance fixation or immobilization to the surface.
- 145. The array of claim 144, wherein said treatment is carried out using an amine or amide compound.
- 146. The array of claim 145, wherein said amine compound is selected from the group consisting of duodecadiamine (DDA), polylysine (PPL), aminopropyltriethoxysilane or a combination of any of the foregoing.
- 147. The array of claim 145, wherein said amide compound comprises formamide.

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- 148. The array of claim 144, wherein said treatment is carried out using a dispersive compound.
- 149. The array of claim 148, wherein said dispersive compound comprises ammonium acetate.
- 150. The array of claim 144, wherein said treatment is carried out using an epoxy compound.
- 151. The array of claim 144, wherein said treatment is carried out using an amine compound and an epoxy compound.
- 152. The array of any of claims 143, 144, 145, 146, 147, 148, 149, 150 or 151, wherein said substrate surface is porous or non-porous.
- 153. The array of claim 152, wherein said porous substrate surface comprises a porous polymeric material.
- 154. The array of claim 153, wherein said polymeric material is selected from the group consisting of dextran, cellulose and nitrocellulose.
- 155. The array of claim 152, wherein said non-porous substrate surface is selected from the group consisting of siliceous matter and non-porous polymeric material.
- 156. The array of claim 155, wherein said siliceous matter comprises glass or a glass-coated surface.
- 157. The array of claim 156, wherein said glass or glass-coated surface comprises porous glass.
- 158. The array of claim 156, wherein said glass or glass-coated surface is selected from the group consisting of wells, depressions, tubes, cuvettes and an apparatus that comprises a plurality of said wells, tubes or cuvettes.

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- 159. The array of claim 158, wherein said wells comprise microtiter wells.
- 160. The array of claim 155, wherein said non-porous polymeric material comprises a plastic.
- 161. The array of claim 160, wherein said plastic is selected from the group consisting of polyethylene, polypropylene, polystyrene and epoxy.
- 162. The array of claim 143, wherein said nucleic acid strands are fixed or immobilized directly or indirectly to said substrate surface.
- 163. The array of claim 143, wherein said nucleic acid strands are single-stranded or double-stranded.
- 164. The array of claim 163, wherein said nucleic acid strands are selected from the group consisting of DNA and RNA, a DNA-RNA hybrid, or combinations thereof.
- 165. The array of claim 164, wherein said nucleic acid strands comprise nucleic acid probe sequences complementary to a target nucleic acid sequence of interest.
- 166. The array of claim 143, wherein said nucleic acid probe sequences are unlabeled.
- 167. The array of claim 163, wherein at least one of said double-stranded nucleic acid strands produces a soluble signal generated or generatable from a chemical label or labels comprising a signaling moiety or moieties.
- 168. The array of claim 167, wherein said label or labels are the signaling moiety or moieties.
- 169. The array of claim 168, wherein the signaling moiety or moieties of said label or labels are directly or indirectly attached thereto.

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- 170. The array of claim 167, wherein said labeled nucleic acid strand comprises a nucleic acid sequence sought to be identified or sequenced.
- 171. The array of claim 163, wherein said label or labels are attached directly or indirectly to one or more nucleotides in said nucleic acid strand.
- 172. The array of claim 163, wherein said label or labels are indirectly attached to one or more nucleotides through the formation of a complex.
- 173. The array of claim 172, wherein said complex is selected from the group consisting of biotin and avidin, biotin and streptavidin, a sugar and a lectin, and an antigen and an antibody.
- 174. The array of claim 167, wherein said label or labels are indirectly attached to one or more nucleotides through a bridging moiety.
- 175. The array of claim 167, wherein said signaling moiety or moieties are selected from the group consisting of an enzyme, a co-enzyme, a chelating agent, a chromagen, a fluorescent agent and a chemiluminescent agent.
- 176. The array of claim 167, wherein said soluble signal is generated or generatable from a chromagen, or by fluorescence or chemiluminescence.
- 177. The array of claim 167, wherein said soluble signal is quantifiable or detectable by a technique selected from the group consisting of photometric techniques and colorimetric techniques.
- 178. The array of claim 177, wherein said photometric techniques comprise spectrophotometric techniques.
- 179. The array of claim 167, wherein said soluble signal is selected from the group consisting of a colored product, a chemiluminescent product and a fluorescent product.

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- 180. An apparatus which comprises the array of any of claims 143 to 151, or 153 to 179, wherein said substrate surface is porous or non-porous.
- 181. A transparent non-porous or translucent non-porous system capable of retaining or containing a fluid or solution, which system comprises the array of any of claims 143 to 151, or 153 to 179, wherein said substrate surface is porous or non-porous.
- 182. The system of claim 181, wherein said substrate surface is contained within the transparent non-porous or translucent non-porous system.

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